

Substitute for form 1449/PTO				<i>Complete if Known</i>	
				Application Number	10/728,665
				Filing Date	December 5, 2003
				First Named Inventor	Satyanarayana MEDICHERLA
				Art Unit	1614
				Examiner Name	A. Marschel
Sheet	1	of	3	Attorney Docket Number	219002032800

U.S. PATENT DOCUMENTS					
Examiner Initials*	Cite No. ¹	Document Number Number-Kind Code ² (if known)	Publication Date MM-DD-YYYY	Name of Patentee or Applicant of Cited Document	Pages, Columns, Lines, Where Relevant Passages or Relevant Figures Appear
FOREIGN PATENT DOCUMENTS					
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NON PATENT LITERATURE DOCUMENTS					
Examiner Initials*	Cite No. ¹	Include name of the author (in CAPITAL LETTERS), title of the article (when appropriate), title of the item (book, magazine, journal, serial, symposium, catalog, etc.), date, page(s), volume-issue number(s), publisher, city and/or country where published.			
	1.	ADAMS et al., "Pyrimidinylimidazole Inhibitors of CSBP/P38 Kinase Demonstrating Decreased Inhibition of Hepatic Cytochrome P450 Enzymes," Bioorg. Med. Chem. Lett. 8:3111-3116 (1998)			
	2.	BADGER et al., "Pharmacological Profile of SB 203580, a Selective Inhibitor of Cytokine Suppressive Binding Protein/p38 Kinase, in Animal Models of Arthritis, Bone Resorption, Endotoxin Shock and Immune Function," J. Pharmacol. Exp. Ther. 279:1453-1461 (1996)			
	3.	BALASA et al., "Islet-Specific Expression of IL-10 Promotes Diabetes in Nonobese Diabetic Mice Independent of Fas, Perforin, TNF Receptor-1, and TNF Receptor-2 Molecules," J. Immunology 165:2841-2849 (2000)			
	4.	BLAIR, "Regulation of Glucose Transport and Glycogen Synthesis in L6 Muscle Cells During Oxidation Stress," J. Biol. Sci. 17:36293-36299 (1999)			
	5.	BURYSEK et al., "The Serine Protease Plasmin Triggers Expression of MCP-1 and CD40 in Human Primary Monocytes via Activation of p38 MAPK and Janus Kinase (JAK)/STAT Signaling Pathways," J. Biol. Sci. 17:36293-36299 (1999)			
	6.	CASTEELS et al., "Prevention of Autoimmune Destruction of Syngeneic Islet Grafts in Spontaneously Diabetic Nonobese Diabetic Mice by a Combination of a Vitamin D ₃ Analog and Cyclosporine," Transplantation 65:1225-1232 (1998)			
	7.	CHEN et al., "Monocyte Chemoattractant Protein-1 is Expressed in Pancreatic Islets from Prediabetic NOD Mice and in Interleukin-1 β -Exposed Human and Rat Islet Cells," Diabetologia 44:325-332 (2001)			
	8.	COLLIS et al., "RPR203494 a Pyrimidine Analogue of the p38 Inhibitor RPR200765A with an Improved In Vitro Potency," Bioorg. Med. Chem. Lett. 11:693-696 (2001)			
	9.	CRUMP, "The Honeymoon Period in Non-Insulin-Dependent Diabetes Mellitus," J. Family Practice 25:78-82 (1987)			
	10.	DEBUSSCHE et al., "Course of Pancreatic β Cell Destruction in Prediabetic NOD Mice: A Histomorphometric Evaluation," Diabète & Métabolisme (Paris) 20:282-290 (1994)			

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11.	DE LASZLO et al., "Pyrroles and Other Heterocycles as Inhibitors of p38 Kinase," <i>Bioorg. Med. Chem. Lett.</i> 8:2698 (1998)	
12.	FIJEN et al., "Suppression of the Clinical and Cytokine Response to Endotoxin by RWJ-67657, a p38 Mitogen-Activated Protein-Kinase Inhibitor, in Healthy Human Volunteers," <i>Clin. Exp. Immunol.</i> 124:16-20 (2001)	
13.	GALLAGHER et al., "Regulation of Stress-Induced Cytokine Production by Pyridinylimidazoles Inhibition of CSBP Kinase," <i>Bioorg. Med. Chem.</i> 5:49-64 (1997)	
14.	GOEBELER et al., "The MKK6/p38 Stress Kinase Cascade Is Critical for Tumor Necrosis Factor- α -Induced Expression of Monocyte-Chemoattractant Protein-1 in Endothelial Cells," <i>Blood</i> 93:857-865 (1999)	
15.	HALE et al., "Differential Expression and Activation of p38 Mitogen-Activated Protein Kinase α , β , γ , and δ in Inflammatory Cell Lineages," <i>J. Immunol.</i> 162:4246-4252 (1999)	
16.	HANCOCK et al., "Suppression of Insulitis in Non-Obese Diabetic (NOD) Mice by Oral Insulin Administration Is Associated with Selective Expression of Interleukin-4 and -10, Transforming Growth Factor- β , and Prostaglandin-E," <i>Am. J. Patho.</i> 147:1194-1199 (1995)	
17.	HEINZE and THON, "Honeymoon Period in Insulin-Dependent Diabetes Mellitus," <i>Pediatrician</i> 12:208-212 (1985)	
18.	HERLAAR and BROWN, "p38 MAPK Signalling Cascades in Inflammatory Disease," <i>Molec. Med. Today</i> 5:439-447 (1999)	
19.	HOSKER and TURNER, "Insulin Treatment of Newly-Presenting Ketotic Diabetic Patients into the Honeymoon Period," <i>Lancet</i> 18:633-635 (1982)	
20.	JACKSON et al., "Pharmacological Effects of SB 220025, a Selective Inhibitor of P38 Mitogen-Activated Protein Kinase, in Angiogenesis and Chronic Inflammatory Disease Models," <i>J. Pharmacol. Exp. Ther.</i> 284:687-692 (1998)	
21.	KEYMEULEN and SOMERS, "Immunointervention in Type 1 (Insulin-Dependent) Diabetes Mellitus," <i>Acta Clinica Belgica</i> 48:86-95 (1993)	
22.	KIMBLE et al., "Simultaneous Block of Interleukin-1 and Tumor Necrosis Factor Is Required to Completely Prevent Bone Loss in the Early Postovariectomy Period," <i>Endocrinol.</i> , 136:3054-3061 (1995)	
23.	KUMAR et al., "Novel Homologues of CSBP/p38 MAP Kinase: Activation, Substrate Specificity and Sensitivity to Inhibition by Pyridinyl Imidazoles," <i>Biochem. Biophys. Res. Comm.</i> 235:533-538 (1997)	
24.	LEE et al., "Inhibition of p38 MAP Kinase as a Therapeutic Strategy," <i>Immunopharmacology</i> 47:185-201 (2000)	
25.	MAHON et al., "Lessons Learned from Use of Cyclosporine for Insulin-Dependent Diabetes Mellitus," <i>Ann. N.Y. Acad. Sci.</i> 696:351-363 (1993)	
26.	MAKINO et al., "Breeding of a Non-Obese, Diabetic Strain of Mice," <i>Exp. Animal</i> 29:1-13 (1980)	
27.	MCLAY et al., "The Discovery of RPR 200765A, p38 MAP Kinase Inhibitor Displaying a Good Oral Anti-Arthritic Efficacy," <i>Bioorg. Med. Chem.</i> 9:537-554 (2001)	
28.	MIYAZAKI et al., "Predominance of T Lymphocytes in Pancreatic Islets and Spleen of Pre-Diabetic Non-Obese Diabetic (NOD) Mice: A Longitudinal Study," <i>Clin. Exp. Immuno.</i> 60:622-630 (1985)	
29.	MORI et al., "Preventive Effects of Cyclosporin on Diabetes in NOD Mice," <i>Diabetologia</i> 29:244-247 (1986)	
30.	PALMER and MCCULLOCH, "Perspective in Diabetes, Prediction and Prevention of IDDM-	

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		1991," Diabetes 40:943-947 (1990)	
31.		PAVLOVIC et al., "Activation of Extracellular Signal-Regulated Kinase (ERK) 1/2 Contributes to Cytokine-Induced Apoptosis in Purified Rat Pancreatic β -Cells," Eur. Cytokine Netw. 11:267-274 (2000)	
32.		POSTE et al., "Lipid Vesicles as Carriers for Introducing Biologically Active Materials into Cells," Meth. Cell Biol. 14:33 (1976)	
33.		RAPOPORT et al., "Thymic T Cell Anergy in Autoimmune Nonobese Diabetic Mice Is Mediated by Deficient T Cell Receptor Regulation of the Pathway of p21ras Activation," J. Exp. Med. 177:1221-1227 (1993)	
34.		RAVELLI, "Macrophage Activation Syndrome," Curr. Opin. Rheumatol. 14:548-552 (2002)	
35.		REVESZ et al., "SAR of 4-Hydroxypiperidine and Hydroxyalkyl Substituted Heterocycles as Novel p38 Map Kinase Inhibitors," Bioorg. Med. Chem. Lett., 10:1261-1264 (2000)	
36.		ROGERS and GIEMBYCZ, "Conquering Airway Inflammation in the 21st Century," Drug Discov. Today 3:532-535 (1998)	
37.		RYU et al., "Reversal of Established Autoimmune Diabetes by Restoration of Endogenous β Cell Function," J. Clin. Invest. 108:63-72 (2001)	
38.		SALOMON et al., "B7/CD28 Costimulation Is Essential for the Homeostasis of the CD4+CD25+ Immunoregulatory T Cells that Control Autoimmune Diabetes," Immunity 12:431-437 (2000)	
39.		SHAPIRO et al., "Combination Therapy with Low Dose Sirolimus and Tacrolimus is Synergistic in Preventing Spontaneous and Recurrent Autoimmune Diabetes in Non-Obese Diabetic Mice," Diabetologia 45:224-230 (2002)	
40.		SPRINKEL et al., "Glucose Potentiates Interleukin-1 β (IL-1 β)-Induced p38 Mitogen-Activated Protein Kinase Activity in Rat Pancreatic Islets of Langerhans," Eur. Cytokine Netw. 12:331-339 (2001)	
41.		SUGIHARA et al., "Ultrastructural and Immunoelectron Microscopic Studies on Infiltrating Mononuclear Cells in Lymphocytic Submandibulitis in NOD Mice," Histol. Histopathol. 4:397-404 (1989)	
42.		TABATABAIE et al., "COX-2 Inhibition Prevents Insulin-Dependent Diabetes in Low-Dose Streptozotocin-Treated Mice," Biochem. Biophys. Res. Comm. 273:699-704 (2000)	
43.		THOMAS and KAY, "Beta Cell Destruction in the Development of Autoimmune Diabetes in the Non-Obese Diabetic (NOD) Mice," Diabetes/Metabolism Res. Rev. 16:251-261 (2000)	
44.		WADSWORTH et al., "RWJ 67657, a Potent, Orally Active Inhibitor of p38 Mitogen-Activated Protein Kinase," J. Pharmacol. Expt. Therapeut. 291:680-687 (1999)	
45.		WANG et al., "Structural Basis of Inhibitor Selectivity in MAP Kinases," Structure 6:1117-1128 (1998)	
46.		YOSHIDA and KIKUTANI, "Genetic and Immunological Basis of Autoimmune Diabetes in the NOD Mouse," Reviews in Immunogenetics 2:140-146 (2000)	
47.		ZHANG et al., "CD28 Co-Stimulation Restores T Cell Responsiveness in NOD Mice by Overcoming Deficiencies in Rac-1/p38 Mitogen-Activated Protein Kinase Signaling and IL-2 and IL-4 Gene Transcription," International Immunology 13:377-384 (2001)	

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